

RAW SEQUENCE LISTING . DATE: 04/09/2001
PATENT APPLICATION: US/09/526,193A TIME: 17:28:37

Input Set : A:\PTO.txt
Output Set: N:\CRF3\04092001\I526193A.raw

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4 <110> APPLICANT: Hayden, Michael R.
5 Brooks-Wilson, Angela R.
6 Pimstone, Simon N.
8 <120> TITLE OF INVENTION: METHODS AND REAGENTS FOR MODULATING
9 CHOLESTEROL LEVELS
11 <130> FILE REFERENCE: 50110/002005
13 <140> CURRENT APPLICATION NUMBER: US 09/526,193A
14 <141> CURRENT FILING DATE: 2000-03-15
16 <150> PRIOR APPLICATION NUMBER: 60/124,702
17 <151> PRIOR FILING DATE: 1999-03-15
19 <150> PRIOR APPLICATION NUMBER: 60/138,048
20 <151> PRIOR FILING DATE: 1999-06-08
22 <150> PRIOR APPLICATION NUMBER: 60/139,600
23 <151> PRIOR FILING DATE: 1999-06-17
25 <150> PRIOR APPLICATION NUMBER: 60/151,977
26 <151> PRIOR FILING DATE: 1999-09-01
28 <160> NUMBER OF SEQ ID NOS: 287
30 <170> SOFTWARE: FastSEQ for Windows Version 4.0
32 <210> SEQ ID NO: 1
33 <211> LENGTH: 2261
34 <212> TYPE: PRT
35 <213> ORGANISM: Homo sapiens
37 <400> SEQUENCE: 1
38 Met Ala Cys Trp Pro Gln Leu Arg Leu Leu Leu Trp Lys Asn Leu Thr
39 1 5 10 15
40 Phe Arg Arg Arg Gln Thr Cys Gln Leu Leu Leu Glu Val Ala Trp Pro
41 20 25 30
42 Leu Phe Ile Phe Leu Ile Leu Ile Ser Val Arg Leu Ser Tyr Pro Pro
43 35 40 45
44 Tyr Glu Gln His Glu Cys His Phe Pro Asn Lys Ala Met Pro Ser Ala
45 50 55 60
46 Gly Thr Leu Pro Trp Val Gln Gly Ile Ile Cys Asn Ala Asn Asn Pro
47 65 70 75 80
48 Cys Phe Arg Tyr Pro Thr Pro Gly Glu Ala Pro Gly Val Val Gly Asn
49 85 90 95
50 Phe Asn Lys Ser Ile Val Ala Arg Leu Phe Ser Asp Ala Arg Arg Leu
51 100 105 110
52 Leu Leu Tyr Ser Gln Lys Asp Thr Ser Met Lys Asp Met Arg Lys Val
53 115 120 125
54 Leu Arg Thr Leu Gln Gln Ile Lys Lys Ser Ser Ser Asn Leu Lys Leu
55 130 135 140
56 Gln Asp Phe Leu Val Asp Asn Glu Thr Phe Ser Gly Phe Leu Tyr His
57 145 150 155 160
58 Asn Leu Ser Leu Pro Lys Ser Thr Val Asp Lys Met Leu Arg Ala Asp
59 165 170 175
60 Val Ile Leu His Lys Val Phe Leu Gln Gly Tyr Gln Leu His Leu Thr
61 180 185 190

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ENTERED

COPY OF
PAPER
NO. 9
WS
3/27/13

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```

62 Ser Leu Cys Asn Gly Ser Lys Ser Glu Glu Met Ile Gln Leu Gly Asp
63      195      200      205
64 Gln Glu Val Ser Glu Leu Cys Gly Leu Pro Arg Glu Lys Leu Ala Ala
65      210      215      220
66 Ala Glu Arg Val Leu Arg Ser Asn Met Asp Ile Leu Lys Pro Ile Leu
67 225      230      235      240
68 Arg Thr Leu Asn Ser Thr Ser Pro Phe Pro Ser Lys Glu Leu Ala Glu
69      245      250      255
70 Ala Thr Lys Thr Leu Leu His Ser Leu Gly Thr Leu Ala Gln Glu Leu
71      260      265      270
72 Phe Ser Met Arg Ser Trp Ser Asp Met Arg Gln Glu Val Met Phe Leu
73      275      280      285
74 Thr Asn Val Asn Ser Ser Ser Ser Thr Gln Ile Tyr Gln Ala Val
75      290      295      300
76 Ser Arg Ile Val Cys Gly His Pro Glu Gly Gly Leu Lys Ile Lys
77 305      310      315      320
78 Ser Leu Asn Trp Tyr Glu Asp Asn Asn Tyr Lys Ala Leu Phe Gly Gly
79      325      330      335
80 Asn Gly Thr Glu Glu Asp Ala Glu Thr Phe Tyr Asp Asn Ser Thr Thr
81      340      345      350
82 Pro Tyr Cys Asn Asp Leu Met Lys Asn Leu Glu Ser Ser Pro Leu Ser
83      355      360      365
84 Arg Ile Ile Trp Lys Ala Leu Lys Pro Leu Leu Val Gly Lys Ile Leu
85      370      375      380
86 Tyr Thr Pro Asp Thr Pro Ala Thr Arg Gln Val Met Ala Glu Val Asn
87 385      390      395      400
88 Lys Thr Phe Gln Glu Leu Ala Val Phe His Asp Leu Glu Gly Met Trp
89      405      410      415
90 Glu Glu Leu Ser Pro Lys Ile Trp Thr Phe Met Glu Asn Ser Gln Glu
91      420      425      430
92 Met Asp Leu Val Arg Met Leu Leu Asp Ser Arg Asp Asn Asp His Phe
93      435      440      445
94 Trp Glu Gln Gln Leu Asp Gly Leu Asp Trp Thr Ala Gln Asp Ile Val
95      450      455      460
96 Ala Phe Leu Ala Lys His Pro Glu Asp Val Gln Ser Ser Asn Gly Ser
97 465      470      475      480
98 Val Tyr Thr Trp Arg Glu Ala Phe Asn Glu Thr Asn Gln Ala Ile Arg
99      485      490      495
100 Thr Ile Ser Arg Phe Met Glu Cys Val Asn Leu Asn Lys Leu Glu Pro
101      500      505      510
102 Ile Ala Thr Glu Val Trp Leu Ile Asn Lys Ser Met Glu Leu Leu Asp
103      515      520      525
104 Glu Arg Lys Phe Trp Ala Gly Ile Val Phe Thr Gly Ile Thr Pro Gly
105      530      535      540
106 Ser Ile Glu Leu Pro His His Val Lys Tyr Lys Ile Arg Met Asp Ile
107 545      550      555      560
108 Asp Asn Val Glu Arg Thr Asn Lys Ile Lys Asp Gly Tyr Trp Asp Pro
109      565      570      575
110 Gly Pro Arg Ala Asp Pro Phe Glu Asp Met Arg Tyr Val Trp Gly Gly

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```

111          580          585          590
112 Phe Ala Tyr Leu Gln Asp Val Val Glu Gln Ala Ile Ile Arg Val Leu
113          595          600          605
114 Thr Gly Thr Glu Lys Lys Thr Gly Val Tyr Met Gln Gln Met Pro Tyr
115          610          615          620
116 Pro Cys Tyr Val Asp Asp Ile Phe Leu Arg Val Met Ser Arg Ser Met
117 625          630          635          640
118 Pro Leu Phe Met Thr Leu Ala Trp Ile Tyr Ser Val Ala Val Ile Ile
119          645          650          655
120 Lys Gly Ile Val Tyr Glu Lys Glu Ala Arg Leu Lys Glu Thr Met Arg
121          660          665          670
122 Ile Met Gly Leu Asp Asn Ser Ile Leu Trp Phe Ser Trp Phe Ile Ser
123          675          680          685
124 Ser Leu Ile Pro Leu Leu Val Ser Ala Gly Leu Leu Val Val Ile Leu
125          690          695          700
126 Lys Leu Gly Asn Leu Leu Pro Tyr Ser Asp Pro Ser Val Val Phe Val
127 705          710          715          720
128 Phe Leu Ser Val Phe Ala Val Val Thr Ile Leu Gln Cys Phe Leu Ile
129          725          730          735
130 Ser Thr Leu Phe Ser Arg Ala Asn Leu Ala Ala Cys Gly Gly Ile
131          740          745          750
132 Ile Tyr Phe Thr Leu Tyr Leu Pro Tyr Val Leu Cys Val Ala Trp Gln
133          755          760          765
134 Asp Tyr Val Gly Phe Thr Leu Lys Ile Phe Ala Ser Leu Leu Ser Pro
135          770          775          780
136 Val Ala Phe Gly Phe Gly Cys Glu Tyr Phe Ala Leu Phe Glu Glu Gln
137 785          790          795          800
138 Gly Ile Gly Val Gln Trp Asp Asn Leu Phe Glu Ser Pro Val Glu Glu
139          805          810          815
140 Asp Gly Phe Asn Leu Thr Thr Ser Val Ser Met Met Leu Phe Asp Thr
141          820          825          830
142 Phe Leu Tyr Gly Val Met Thr Trp Tyr Ile Glu Ala Val Phe Pro Gly
143          835          840          845
144 Gln Tyr Gly Ile Pro Arg Pro Trp Tyr Phe Pro Cys Thr Lys Ser Tyr
145          850          855          860
146 Trp Phe Gly Glu Glu Ser Asp Glu Lys Ser His Pro Gly Ser Asn Gln
147 865          870          875          880
148 Lys Arg Ile Ser Glu Ile Cys Met Glu Glu Glu Pro Thr His Leu Lys
149          885          890          895
150 Leu Gly Val Ser Ile Gln Asn Leu Val Lys Val Tyr Arg Asp Gly Met
151          900          905          910
152 Lys Val Ala Val Asp Gly Leu Ala Leu Asn Phe Tyr Glu Gly Gln Ile
153          915          920          925
154 Thr Ser Phe Leu Gly His Asn Gly Ala Gly Lys Thr Thr Thr Met Ser
155          930          935          940
156 Ile Leu Thr Gly Leu Phe Pro Pro Thr Ser Gly Thr Ala Tyr Ile Leu
157 945          950          955          960
158 Gly Lys Asp Ile Arg Ser Glu Met Ser Thr Ile Arg Gln Asn Leu Gly
159          965          970          975

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```

160 Val Cys Pro Gln His Asn Val Leu Phe Asp Met Leu Thr Val Glu Glu
161          980          985          990
162 His Ile Trp Phe Tyr Ala Arg Leu Lys Gly Leu Ser Glu Lys His Val
163          995          1000          1005
164 Lys Ala Glu Met Glu Gln Met Ala Leu Asp Val Gly Leu Pro Ser Ser
165          1010          1015          1020
166 Lys Leu Lys Ser Lys Thr Ser Gln Leu Ser Gly Gly Met Gln Arg Lys
167 1025          1030          1035          1040
168 Leu Ser Val Ala Leu Ala Phe Val Gly Gly Ser Lys Val Val Ile Leu
169          1045          1050          1055
170 Asp Glu Pro Thr Ala Gly Val Asp Pro Tyr Ser Arg Arg Gly Ile Trp
171          1060          1065          1070
172 Glu Leu Leu Leu Lys Tyr Arg Gln Gly Arg Thr Ile Ile Leu Ser Thr
173          1075          1080          1085
174 His His Met Asp Glu Ala Asp Val Leu Gly Asp Arg Ile Ala Ile Ile
175          1090          1095          1100
176 Ser His Gly Lys Leu Cys Cys Val Gly Ser Ser Leu Phe Leu Lys Asn
177 1105          1110          1115          1120
178 Gln Leu Gly Thr Gly Tyr Tyr Leu Thr Leu Val Lys Lys Asp Val Glu
179          1125          1130          1135
180 Ser Ser Leu Ser Ser Cys Arg Asn Ser Ser Ser Thr Val Ser Tyr Leu
181          1140          1145          1150
182 Lys Lys Glu Asp Ser Val Ser Gln Ser Ser Ser Asp Ala Gly Leu Gly
183          1155          1160          1165
184 Ser Asp His Glu Ser Asp Thr Leu Thr Ile Asp Val Ser Ala Ile Ser
185          1170          1175          1180
186 Asn Leu Ile Arg Lys His Val Ser Glu Ala Arg Leu Val Glu Asp Ile
187 1185          1190          1195          1200
188 Gly His Glu Leu Thr Tyr Val Leu Pro Tyr Glu Ala Ala Lys Glu Gly
189          1205          1210          1215
190 Ala Phe Val Glu Leu Phe His Glu Ile Asp Asp Arg Leu Ser Asp Leu
191          1220          1225          1230
192 Gly Ile Ser Ser Tyr Gly Ile Ser Glu Thr Thr Leu Glu Glu Ile Phe
193          1235          1240          1245
194 Leu Lys Val Ala Glu Glu Ser Gly Val Asp Ala Glu Thr Ser Asp Gly
195          1250          1255          1260
196 Thr Leu Pro Ala Arg Arg Asn Arg Arg Ala Phe Gly Asp Lys Gln Ser
197 1265          1270          1275          1280
198 Cys Leu Arg Pro Phe Thr Glu Asp Asp Ala Ala Asp Pro Asn Asp Ser
199          1285          1290          1295
200 Asp Ile Asp Pro Glu Ser Arg Glu Thr Asp Leu Leu Ser Gly Met Asp
201          1300          1305          1310
202 Gly Lys Gly Ser Tyr Gln Val Lys Gly Trp Lys Leu Thr Gln Gln Gln
203          1315          1320          1325
204 Phe Val Ala Leu Leu Trp Lys Arg Leu Leu Ile Ala Arg Arg Ser Arg
205          1330          1335          1340
206 Lys Gly Phe Phe Ala Gln Ile Val Leu Pro Ala Val Phe Val Cys Ile
207 1345          1350          1355          1360
208 Ala Leu Val Phe Ser Leu Ile Val Pro Pro Phe Gly Lys Tyr Pro Ser

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```

209          1365          1370          1375
210 Leu Glu Leu Gln Pro Trp Met Tyr Asn Glu Gln Tyr Thr Phe Val Ser
211          1380          1385          1390
212 Asn Asp Ala Pro Glu Asp Thr Gly Thr Leu Glu Leu Leu Asn Ala Leu
213          1395          1400          1405
214 Thr Lys Asp Pro Gly Phe Gly Thr Arg Cys Met Glu Gly Asn Pro Ile
215          1410          1415          1420
216 Pro Asp Thr Pro Cys Gln Ala Gly Glu Glu Glu Trp Thr Thr Ala Pro
217 1425          1430          1435          1440
218 Val Pro Gln Thr Ile Met Asp Leu Phe Gln Asn Gly Asn Trp Thr Met
219          1445          1450          1455
220 Gln Asn Pro Ser Pro Ala Cys Gln Cys Ser Ser Asp Lys Ile Lys Lys
221          1460          1465          1470
222 Met Leu Pro Val Cys Pro Pro Gly Ala Gly Gly Leu Pro Pro Pro Gln
223          1475          1480          1485
224 Arg Lys Gln Asn Thr Ala Asp Ile Leu Gln Asp Leu Thr Gly Arg Asn
225          1490          1495          1500
226 Ile Ser Asp Tyr Leu Val Lys Thr Tyr Val Gln Ile Ile Ala Lys Ser
227 1505          1510          1515          1520
228 Leu Lys Asn Lys Ile Trp Val Asn Glu Phe Arg Tyr Gly Gly Phe Ser
229          1525          1530          1535
230 Leu Gly Val Ser Asn Thr Gln Ala Leu Pro Pro Ser Gln Glu Val Asn
231          1540          1545          1550
232 Asp Ala Ile Lys Gln Met Lys Lys His Leu Lys Leu Ala Lys Asp Ser
233          1555          1560          1565
234 Ser Ala Asp Arg Phe Leu Asn Ser Leu Gly Arg Phe Met Thr Gly Leu
235          1570          1575          1580
236 Asp Thr Arg Asn Asn Val Lys Val Trp Phe Asn Asn Lys Gly Trp His
237 1585          1590          1595          1600
238 Ala Ile Ser Ser Phe Leu Asn Val Ile Asn Asn Ala Ile Leu Arg Ala
239          1605          1610          1615
240 Asn Leu Gln Lys Gly Glu Asn Pro Ser His Tyr Gly Ile Thr Ala Phe
241          1620          1625          1630
242 Asn His Pro Leu Asn Leu Thr Lys Gln Gln Leu Ser Glu Val Ala Leu
243          1635          1640          1645
244 Met Thr Thr Ser Val Asp Val Leu Val Ser Ile Cys Val Ile Phe Ala
245          1650          1655          1660
246 Met Ser Phe Val Pro Ala Ser Phe Val Val Phe Leu Ile Gln Glu Arg
247 1665          1670          1675          1680
248 Val Ser Lys Ala Lys His Leu Gln Phe Ile Ser Gly Val Lys Pro Val
249          1685          1690          1695
250 Ile Tyr Trp Leu Ser Asn Phe Val Trp Asp Met Cys Asn Tyr Val Val
251          1700          1705          1710
252 Pro Ala Thr Leu Val Ile Ile Ile Phe Ile Cys Phe Gln Gln Lys Ser
253          1715          1720          1725
254 Tyr Val Ser Ser Thr Asn Leu Pro Val Leu Ala Leu Leu Leu Leu
255          1730          1735          1740
256 Tyr Gly Trp Ser Ile Thr Pro Leu Met Tyr Pro Ala Ser Phe Val Phe
257 1745          1750          1755          1760

```

Please Note:

Use of n and/or Xaa have been detected in the Sequence Listing. Please review the Sequence Listing to ensure that a corresponding explanation is presented in the <220> to <223> fields of each sequence which presents at least one n or Xaa.

VERIFICATION SUMMARY

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Input Set : A:\PTO.txt

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L:730 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:14
L:1004 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:19
L:1524 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:21
L:2083 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:29
L:2084 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:29
L:2085 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:29
L:2259 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:39
L:2400 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:54
L:2469 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:61
L:2546 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:69